Morphologic and functional modifications of cells reacting to changes (physiologic) of the external environment

ATROPHY HYPERTROPHY

HYPERPLASIA

METAPLASIA

From Physiological to pathological





External or internal triggers force the cell toward adaptation, thus modifying proliferation and morphology

- Cell size
- Cell differentiation
- Mitotic rate



Modalities of cellular adaptation

- atrophy = decreased cell size
- *hypertrophy* = increased cell size
- *hyperplasia* = incresed number of cells
- *metaplasia* = change in cell phenotype/function

ATROPHY = Decreased volume of an organ or a cell



PATHOGENESIS: reduced consumption of aminoacid and O₂, reduced protein synthesis, increased protein cabolism

Atrophy: microscopic features

- Decreased cell size
- Tightening of nuclei
- Cytoplasms pigments "wear-and-tear"
- Substitution with adipose or fibrous tissue (hyalinosis)
- Organ shape is preserved, size is decreased and consistency is increased
- Lack of substitution with fibrous or adipose tissue = reduced size

Examples of atrophy

- Lymphatic tissue, bone marrow in elderly people
- Adrenal glands after prolonged steroid therapy
- Inactive scheletal muscle
- Absence of endocrine stimulation on target organs

Dd (differential diagnosis):

Hypoplasia – incomplete growth Agenesia – lack of development of an organ











HYPERPLASIA / HYPERTROPHY



Hypertrophy and hyperplasia

- Respond to well-defined stimuli
- Limited in space and time
- Specific purpose
- Preservation of normal architectural, cytological and functional features of original tissues
- Don't progress
- Regress when trigger disappears

Hypertrophy

Increased cell/organ size

• Physiologic or pathologic

- CAUSES:
 - increased functional requests
 - specific hormonal stimulation (correlative-endocrine)

Hypertrophy

PHYSIOLOGIC: - Pregnant uterus

- Scheletal muscle (Increased muscle activity)

PATHOLOGIC: - Myocardial Hypertrophy (hypertension) - Smooth muscle of hollow organs (stomach, bladder, ureters) (following obstruction)



HYPERPLASIA: Volumetric increase of an organ or part of it, due to increased cell **<u>number</u> —** labile and stable tissue



Pathogenesis of HYPERPLASIA:

- Production of substances which stimulate or potentiate cell growth (*hormones, growth factors, ecc*.)
- Loss of calons (mitosis-inhibitory substances)
- Prolonged cell survival

Histological features of Hyperplasia:

- Increased cell number
- Cell crowding,
- Multistratification
- Increased mitotic rate
- Light (reactive) atypia

ATYPIC HYPERPLASIA **BORDERLINE**" LESION

HYPERPLASIA Reactive alteration of cell Proliferation = Increased number of normal components



BASAL CELL HYPERPLASIA







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GOBLET CELL HYPERPLASIA







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Metaplasia

- Pathologic process of functional differentiation
- Cells adapt to environmental and functional changes, modifying their morphologic appearance
- Transformation of tissue in another type of the same embryologic derivation
- The transformation of a tissue of ectodermic origin into one of mesodermic origin or viceversa **IS IMPOSSIBLE**

METAPLASIA



Adaptive change of cell differentiation = changes of functional specialization





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Epithelial Metaplasia

- **Respiratory epithelium** \longrightarrow Squamous epithelium
- **Mammary gland** \longrightarrow Apocrine cell metaplasia
- **Thyroid** → Oncocytic metaplasia
- **Salivary Glands** \longrightarrow Oncocytic metaplasia









SQUAMOUS METAPLASIA









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Squamous metaplasia of bronchial epithelium







Intestinal metaplasia of stomach







Dysplasia

Pathologic process of differentiation (maturation)

Spontaneously regresses with withdrawal of the cause

- Organ (e.g., renal) dysplasia = altered maturation during developmental stages
- Cell dysplasia = maturative arrest before endstage differentiation

Cell Dysplasia

- Maturative arrest
- Expansion of the proliferative compartment
- Altered architecture, irregular stratification, > thickness
- Cytoplasmic and nuclear modifications (increased N/C ratio)
- Sites: skin (Bowen's dis.), oesophagus (Barrett's), stomach and colon (adenomatous polyps, IBD), cervix



Altered cell maturation = differentiation arrest with intact proliferative activity





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Reactive atypia

Morphological features

- Intercellular cohesion (clusters)
- Nuclear polarization
- Terminal plaque and cilia
- Nuclear hyperchomasia, nucleoli
- Smooth nuclear contour
- Multinucleation
- Lack of mitotic activity











